

<120> CLOSTRIDIAL TOXIN DERIVATIVES AND METHODS FOR TREATING PAIN

<130> botulinum-subP/pain/D2875

<140>09/489,667

<141>2000-01-19

<160> 14

<170> PatentIn Ver. 2.1

<210>1

<211>11

<212> PRT

<213> Unknown Organism

<220>

<221> MOD_RES

<222>(11)

<223> AMIDATION

<220>

<223> Description of Unknown Organism: This fragment is substance P and is very well known in the art.

<220>

<223> The Met at position 11 is Met-amide.

<300>

<310> 5891842

<311>1996-04-12

<312> 1999-04-06

<400> 1

Arg Pro Lys Pro Gln Gln Phe Phe Gly Leu Met

1

5

10

```
<210>2
<211>12
<212> PRT
<213> Unknown Organism
<220>
<223> Description of Unknown Organism: Precursor to
   substance P, which is very well known in the art.
<300>
<310> 5891842
<311> 1996-04-12
<312> 1999-04-06
<300>
<301> Shimonka,
   et al.,
<303> J. Neurochem.
<304> 59
<306> 81-92
<307> 1992
<400> 2
Arg Pro Lys Pro Gln Gln Phe Phe Gly Leu Met Gly
 1
          5
                     10
<210>3
<211>13
<212> PRT
<213> Unknown Organism
<220>
<223> Description of Unknown Organism: This fragment is
   a precursor to substance P and is very well known
   in the art.
<300>
<310> 5891842
<311>1996-04-12
<312> 1999-04-06
```

<300>

```
<301> Shimonka,
   et al.,
<303> J. Neurochem.
<304> 59
<306> 81-92
<307> 1992
<400>3
Arg Pro Lys Pro Gln Gln Phe Phe Gly Leu Met Gly Lys
          5
                      10
<210>4
<211>14
<212> PRT
<213> Unknown Organism
<220>
<223> Description of Unknown Organism: This fragment is a
   precursor to substance P and is very well known in
   the art.
<300>
<310> 5891842
<311> 1996-04-12
<312> 1999-04-06
<300>
<301> Shimonka,
   et al.,
<303> J. Neurochem.
<304> 59
<306> 81-92
<307> 1992
<400>4
Arg Pro Lys Pro Gln Gln Phe Phe Gly Leu Met Gly Lys Arg
          5
                      10
<210>5
<211>12
<212> PRT
```

```
<213> Artificial Sequence
<220>
<223> Description of Artificial Sequence: This fragment
   is a carboxy-ester synthetic precursor to
   substance P.
<220>
<223> The Gly at the carboxy terminal (Gly at position
   12) is methylated.
<300>
<310> 5891842
<311> 1996-04-12
<312> 1999-04-06
<300>
<301> Lee,
   et al.,
<303> Eur. J. Biochem.
<304>114
<306> 315-327
<307> 1981
<300>
<301> Pernow, B.
<303> Pharmacol. Rev.
<304>35
<306> 86-138
<307>1983
<300>
<301> Regoli,
   et al.,
<303> TIPS
<304>9
<306> 290-295
<307> 1988
<400> 5
Arg Pro Lys Pro Gln Gln Phe Phe Gly Leu Met Gly
 1
```

10

```
<210>6
<211>13
<212> PRT
<213> Artificial Sequence
<220>
<223> Description of Artificial Sequence: This is a
   carboxy ester synthetic precursor to substance P.
<220>
<223> The Lys at the carboxy-terminus (Lys at position
   13) is methylated.
<300>
<310> 5891842
<311>1996-04-12
<312> 1999-04-06
<300>
<301> Lee,
   et al.,
<303> Eur. J. Biochem.
<304>114
<306> 315-327
<307> 1981
<300>
<301> Pernow, B.
<303> Pharmacol. Rev.
<304> 35
<306> 86-138
<307> 1983
<300>
<301> Regoli,
   et al.,
<303> TIPS
<304>9
<306> 290-295
<307>1988
<400>6
Arg Pro Lys Pro Gln Gln Phe Phe Gly Leu Met Gly Lys
```

1 5 10

```
<210>7
<211>14
<212> PRT
<213> Artificial Sequence
<220>
<223> Description of Artificial Sequence: This is a
   carboxy ester sythetic precursor to substance P.
<220>
<223> The Arg at the carboxy-terminus (Arg at position
   14) is methylated.
<300>
<310> 5891842
<311> 1996-04-12
<312> 1999-04-06
<300>
<301> Lee,
   et al.,
<303> Eur. J. Biochem.
<304>114
<306> 315-327
<307> 1981
<300>
<301> Pernow, B.
<303> Pharmacol. Rev.
<304> 35
<306> 86-138
<307> 1983
<300>
<301> Regoli,
```

et al., <303> TIPS <304> 9

<306> 290-295 <307> 1988

```
<400> 7
Arg Pro Lys Pro Gln Gln Phe Phe Gly Leu Met Gly Lys Arg
 1
           5
                      10
<210>8
<211>12
<212> PRT
<213> Artificial Sequence
<220>
<223> Description of Artificial Sequence: This is a
   carboxy ester synthetic precursor to substance P.
<220>
<223> The Gly at the carboxy terminal (Gly at position
   12) is ethylated.
<300>
<310> 5891842
<311> 1996-04-12
<312> 1999-04-06
<300>
<301> Lee,
   et al.,
<303> Eur. J. Biochem.
<304>114
<306> 315-327
<307>1981
<300>
<301> Pernow, B.
<303> Pharmacol. Rev.
<304> 35
<306> 86-138
<307> 1983
<300>
<301> Regoli,
   et al.,
<303> TIPS
<304>9
```

```
<306> 290-295
<307> 1988
<400>8
Arg Pro Lys Pro Gln Gln Phe Phe Gly Leu Met Gly
          5
 1
                      10
<210>9
<211>13
<212> PRT
<213> Artificial Sequence
<220>
<223> Description of Artificial Sequence: This is a
   carboxy ester synthetic precursor to substance P.
<220>
<223> The Lys at the carboxy terminal (Lys at position
   13) is ethylated.
<300>
<310> 5891842
<311> 1996-04-12
<312> 1999-04-06
<300>
<301> Lee,
   et al.,
<303> Eur. J. Biochem.
<304> 114
<306> 315-327
<307>1981
<300>
<301> Pernow, B.
<303> Pharmacol. Rev.
<304>35
<306> 86-138
<307> 1983
<300>
<301> Regoli,
```

```
et al.,
<303> TIPS
<304>9
<306> 290-295
<307>1988
<400>9
Arg Pro Lys Pro Gln Gln Phe Phe Gly Leu Met Gly Lys
 1
           5
                      10
<210>10
<211>14
<212> PRT
<213> Artificial Sequence
<220>
<223> Description of Artificial Sequence: This is a
   carboxy ester synthetic precursor to substance P.
<220>
<223> The Arg at the carboxy terminal (Arg at position
   14) is ethylated.
<300>
<310> 5891842
<311>1996-04-12
<312> 1999-04-06
<300>
<301> Lee,
   et al.,
<303> Eur. J. Biochem.
<304> 114
<306> 315-327
<307> 1981
<300>
<301> Pernow, B.
<303> Pharmacol. Rev.
<304>35
<306> 86-138
<307> 1983
```

```
<300>
<301> Regoli,
   et al.,
<303> TIPS
<304>9
<306> 290-295
<307> 1988
<400>10
Arg Pro Lys Pro Gln Gln Phe Phe Gly Leu Met Gly Lys Arg
                      10
<210>11
<211>4
<212> PRT
<213> Unknown Organism
<220>
<223> This sequence is made up by the first four amino
   acids of substance P.
<220>
<223> Description of Unknown Organism: This is a
   naturally occuring amino terminal peptide fragment
   derived from substance P.
<300>
<310> 5891842
<311>1996-04-12
<312> 1999-04-06
<300>
<301> Stewart,
   et al.,
<303> Nature
<304> 262
<306> 784-785
<307> 1986
<300>
<301> Skilling,
   et al.,
```

```
<303> J. Neurosci.
<304>10
<306> 309-1318
<307> 1990
<400>11
Arg Pro Lys Pro
 1
<210>12
<211>7
<212> PRT
<213> Unknown Organism
<220>
<223> Description of Unknown Organism: This is a
   naturally occuring amino terminal peptide fragment
   derived from substance P.
<220>
<223> This fragment is made up of the first seven amino
   acids of substance P.
<300>
<310> 5891842
<311>1996-04-12
<312> 1999-04-06
<300>
<301> Stewart,
   et al.,
<303> Nature
<304> 262
<306> 784-785
<307>1986
<300>
<301> Skilling,
   et al.,
<303> J. Neurosci.
<304> 10
<306> 309-1318
```

```
<307> 1990
<300>
<301> Lavielle,
   et al.,
<303> Biochem. Pharmacol.
<304>37
<306>41-
<307> 1988
<400> 12
Arg Pro Lys Pro Gln Gln Phe
           5
<210> 13
<211>9
<212> PRT
<213> Unknown Organism
<220>
<223> Description of Unknown Organism: This is a
   naturally occuring amino terminal peptide fragment
   derived from substance P.
<220>
<223> This fragment is made up of the first nine amino
   acids of substance P.
<300>
<310> 5891842
<311> 1996-04-12
<312> 1999-04-06
<300>
<301> Stewart,
   et al.,
<303> Nature
<304> 262
<306> 784-785
<307> 1986
<300>
```

```
<301> Skilling,
   et al.,
<303> J. Neurosci.
<304>10
<306> 309-1318
<307> 1990
<400> 13
Arg Pro Lys Pro Gln Gln Phe Phe Gly
<210> 14
<211>11
<212> PRT
<213> Artificial Sequence
<220>
<223> Description of Artificial Sequence: This is an
   analog of substance P. This analog contains
   disulfide Cys-Cys bridges.
<223> The Cys at position 3 bridges with the Cys at
   position 6.
<220>
<221> MOD_RES
<222>(11)
<223> AMIDATION
<220>
<223> The Met at position 11 is Met-amide.
<300>
<310> 5891842
<311>1996-04-12
<312> 1999-04-06
<300>
<301> Lavielle,
   et al.,
<303> Biochem. Pharmacol.
```